

1	From the INTERNATIONAL BUREAU
PCT	То:
NOTIFICATION OF ELECTION  (PCT Rule 61.2)	Assistant Commissioner for Patents United States Patent and Trademark Office Box PCT Washington, D.C.20231 ETATS-UNIS D'AMERIQUE
Date of mailing (day/month/year) 23 May 2000 (23.05.00)	in its capacity as elected Office
International application No. PCT/US99/20308	Applicant's or agent's file reference
International filing date (day/month/year) 25 August 1999 (25.08.99)	Priority date (day/month/year) 27 August 1998 (27.08.98)
MESSING, Joachim et al	
1. The designated Office is hereby notified of its election made.    X   in the demand filed with the International Preliminal 20 March 200	ry Examining Authority on: 10 (20.03.00)  rnational Bureau on:
The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland	Authorized officer  C. Cupello

Facsimile No.: (41-22) 740.14.35

Telephone No.: (41-22) 338.83.38

International application No.
PCT/US99/20308

A. CLA	SSIFICATION OF SUBJECT MATTER				
` '	Please See Extra Sheet.				
	Please See Extra Sheet.		.:e:	00	
<del></del>	o International Patent Classification (IPC) or to both	nadonal clas	Sincation and II	PC	
	DS SEARCHED				
Minimum d	ocumentation searched (classification system followe	d by classific	cation symbols)		
<b>U.S.</b> :	435/69.1, 320.1, 410, 412, 419, 468; 536/23.6; 800/	278, 287, 29:	5, 298, 320.1		-
Documentat	ion searched other than minimum documentation to the	e extent that s	such documents	are included	in the fields searched
	ata base consulted during the international search (na gricola, Biosis, Caplus, EMBL, Genseq, EST	ame of data b	pase and, where	practicable,	search terms used)
C. DOC	UMENTS CONSIDERED TO BE RELEVANT	-			
Category*	Citation of document, with indication, where ap	propriate, of	the relevant pas	sages	Relevant to claim No.
Y	KIRIHARA et al. Isolation And Seque Methionine-Rich 10-kDa Zein Protein Vol. 71, pages 359-370, especially page	From N	laize. Gene	. 1988,	1-21
Y	CHAUDHURI et al. Allele-specific Pa Posttranscriptional Regulator Of Zein Acad. Sci, USA. May 1994, Vol. 91, pages 4868-4870.	Accumu	lation. Proc	c. Natl.	1-18
Y	CRUZ-ALVAREZ et al. Post-trai Methionine Content In Maize Kernels. 225, pages 331-339, especially pages 3	Mol. Gen	. Genet. 199	91, Vol.	1-18
X Furth	er documents are listed in the continuation of Box C	. 🔲	See patent famil	ly annex.	
"A" doc	cial categories of cited documents:	dat		t with the appli	mational filing date or priority cation but cited to understand invention
"B" eari	the of particular relevance  tier document published on or after the international filing date  tument which may throw doubts on priority claim(s) or which is  do to establish the publication date of another citation or other	co: wb		nnot be consider	claimed invention cannot be ed to involve an inventive step
spe	cial reason (as specified)  cument referring to an oral disclosure, use, exhibition or other	COI	nsidered to involve mbined with one or s	en inventive more other such	claimed invention cannot be step when the document is documents, such combination
"P" doc	nument published prior to the international filing date but later than priority date claimed		ing obvious to a pen cument member of t		
	actual completion of the international search	Date of mai	ling of the inter	mational sear	rch report
24 NOVE	MBER 1999	07 D	EC 1999		
Commission Box PCT	nailing address of the ISA/US ner of Patents and Trademarks	Authorized		M.C.	10
	, D.C. 20231 o. (703) 305-3230	Telephone 1	-1	7101VV	X/
PACAMBLE N	u	LICIEDIIONE I	MAR. 1 / (1941 141)	10-0150	- ·

International application No.
PCT/US99/20308

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No
ť	SWARUP et al. Determinants Of The High-methionine Trait In Wild And Exotic Germplasm May Have Escaped Selection During Early Cultivation Of Maize. The Plant Journal. 1995, Vol. 8, No. 3, pages 359-368, especially pages 361-364.	1-18
7	PIETRZAK et al. Expression In Plants Of Two Bacterial Antibiotic Resistance Genes After Protoplast Transformation With A New Plant Expression Vector. Nucl. Acids Res. 1986, Vol. 14, No. 14, pages 5857-5868, especially pages 5858, 5859, 5861 and 5862.	15

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Box i Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This international report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
2. Claims Nos.: because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
Please See Extra Sheet.
1. X As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:
4. No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark on Protest The additional search fees were accompanied by the applicant's protest.
X No protest accompanied the payment of additional search fees.

International application No. PCT/US99/20308

A. CLASSIFICATION OF SUBJECT MATTER: IPC (6):

C12N 5/04, 15/29, 15/09, 15/11, 15/63, 15/64, 15/66, 15/67, 15/82; A01H 5/00, 5/10

A. CLASSIFICATION OF SUBJECT MATTER: US CL :

435/69.1, 320.1, 410, 412, 419, 468; 536/23.6; 800/278, 287, 295, 298, 320.1

BOX II. OBSERVATIONS WHERE UNITY OF INVENTION WAS LACKING This ISA found multiple inventions as follows:

This application contains the following inventions or groups of inventions which are not so linked as to form a single inventive concept under PCT Rule 13.1. In order for all inventions to be searched, the appropriate additional search fees must be paid.

Group I, claims 1-18, drawn to a first product, a DNA construct encoding a delta-zein, linked to a promoter and a 3' untranslated region (UTR) modified to be devoid of dzrl binding sites, and vectors and plants transformed with said DNA construct, and a first method, comprising making high methionine com with said DNA construct. Group II, claim(s) 19-21, drawn to a second product, an isolated nucleic acid comprising a 3' UTR of a 10 kDa zein gene, and a chimeric gene comprising any coding sequence operably linked to said UTR. The inventions listed as Groups I and II do not relate to a single inventive concept under PCT Rule 13.1 because, under PCT Rule 13.2, they lack the same or corresponding special technical features for the following reasons: The modified 3' UTR of the first product in Group I is not shared with the non-modified 3' UTR of the second product in Group II. The non-modified UTR of Group II also is not shared or used by the method of Group I. Further, the sequences of both groups may be obtained by alternative means, such as chemical synthesis.

## PATENT COOPERATION TREATY

# **PCT**

REC'D	1	1	JAN	2001
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WPO

PCT

## INTERNATIONAL PRELIMINARY EXAMINATION REPORT

(PCT Article 36 and Rule 70)

15

Applicant's or agent's file reference RUT 99-0002P	FOR FURTHER ACTION	See Notification of Transmittal of Internation Preliminary Examination Report (Form PCT/IPEA/41)
International application No.	International filing date (day/n	month/year) Priority date (day/month/year)
PCT/US99/20308	25 AUGUST 1999	27 AUGUST 1998
International Patent Classification (IPC) Please See Supplemental Sheet.	or national classification and IF	PC
Applicant RUTGERS, THE STATE UNIVERSI	TY OF NEW JERSEY	
Examining Authority and is  2. This REPORT consists of a  This report is also accombeen amended and are the	total of sheets.  sheets.  spanied by ANNEXES, i.e., sheet basis for this report and/or sheetin 607 of the Administrative	ets of the description, claims and/or drawings which ha neets containing rectifications made before this Authori
3. This report contains indication		tems:
I X Basis of the repo	·Ft	
II Priority		
III Non-establishmer	nt of report with regard to no	ovelty, inventive step or industrial applicability
IV X Lack of unity of	invention	
	nt under Article 35(2) with reg anations supporting such statem	gard to novelty, inventive step or industrial applicabili- ment
VI Certain documents	cited	
VII Certain defects in t	the international application	
VIII X Certain observation	ns on the international applicati	ion
Date of submission of the demand	Date	e of completion of this report
20 MARCH 2000	2	20 NOVEMBER 2000
Name and mailing address of the IPEA	ŧ.	norized officer  DELLA MAE COLLINS
Commissioner of Patents and Trader Box PCT Washington, D.C. 20231		ASHWIN MEHTA PARALEGAL SPECIALIS
Facsimile No. (703) 305-3230	Teler	phone No. (703) 308-0196 TECHNOLOGY CENTER 16

International application No.

PCT/US99/20308

I.	Basis o	of the report		
1. V	Vith rega	rd to the elements of the intern	national application:*	
_		international application as		
	☱	description:	5	
L	Δ	es <u>1-34</u>		as originally filed
		es NONE		
	_		, filed with the letter of	
_	<b>-</b>		<del></del>	
Ŀ	<u></u>	claims: as 35-37		
			an amonded (as set) as with	, as originally filed
		es NONE NONE	, as amended (together with	
			, filed with the letter of	
	F0		, ,	
Г	X the	drawings:		
_	page	es1-8		, as originally filed
		s NONE		, filed with the demand
	page	s NONE	, filed with the letter of	
_	<b>.</b> .			
		sequence listing part of the ces		
	page	NONE	, filed with the letter of	, filed with the demand
	Pub		, med with the letter of	
	the l	anguage of publication of anguage of the translation fun	the international application (under Rule 48.3 nished for the purposes of international preliminary	(b)).
_ [	With rega	ard to any <b>nucleotide and/o</b> ary examination was carried	r amino acid sequence disclosed in the internat	ional application, the international
<u> </u>	≌ conta	ained in the international a	pplication in printed form.	
	filed	together with the internati	ional application in computer readable form.	
Γ	furni	shed subsequently to this	Authority in written form.	
Ē	 furni	shed subsequently to this	Authority in computer readable form.	
  -	르		ntly furnished written sequence listing does not	no howard the direct own in the
L	⊔ interi —	national application as filed	has been furnished.	,
L	The s	statement that the information furnished.	recorded in computer readable form is identical t	o the writen sequence listing has
4.	The	amendments have resulted	in the cancellation of:	
	X	the description, pages	NONE	
	X	the claims, Nos.	NONE	
	$\mathbf{x}$	the drawings, sheets/fig	NONE	
5. <b>Г</b>	This		some of) the amendments had not been made, since	a they have been considered to ac
· L			indicated in the Supplemental Box (Rule 70.2(c)).*	
ın	eplaceme	nt sheets which have been furn port as "originally filed" and	ished to the receiving Office in response to an invita- are not annexed to this report since they do not	tion under Article 14 are referred to
**A	ny repla	cement sheet containing such	n amendments must be referred to under item 1 a	and annexed to this report.

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IV	7. Lack of unity of invention	
1.	In response to the invitation to restrict or pay additional fees the applicant has:	1
	restricted the claims.	
!	X paid additional fees.	
	paid additional fees under protest.	
	neither restricted nor paid additional fees.	
2.	This Authority found that the requirement of unity of invention is not complied with and chose, according to Rul not to invite the applicant to restrict or pay additional fees.	68.1,
3.	This Authority considers that the requirement of unity of invention in accordance with Rules 13.1, 13.2 and 13.3 is	1
	complied with.	
	X not complied with for the following reasons:	ļ
1	Please See Supplemental Sheet.	
4.	Consequently, the following parts of the international application were the subject of international preliminary examination in establishing this report:	
	X all parts.	
	the parts relating to claims Nos	

International application No.

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Inventive Step (IS)  Claims 1-18, 20, and 21  Claims 19  Industrial Applicability (IA)  Claims 1-21  Claims NONE  Claims 1-21  Claims 19 lacks novelty under PCT Article 33(2) as being anticipated by Kirihara et al.  The claim is broadly drawn towards any isolated nucleic acid comprising the 3' UTR of any 10 kDa zein gene. Kirihara et al teach the nucleotide sequenc3e of the 10 kDa zein gene from maize. Sequences found within tuntranslated region in the 3' terminus are shown (page 362).  Claim 19 lacks an inventive step under PCT Article 33(3) as being obvious over Kirihara et al.  Kirihara et al is described above. The claims lack an inventive step as the reference describes a product that is encompassed by the claims.  Claims 1-18, 20, and 21 meet the criteria set out in PCT Article 33(2) and (3), because the prior art does not teach or fa suggest a DNA construct comprising the coding sequences of a delta-zein operably linked to a 3' UTR that has been modified to no longer contain binding sites for the dzr1 negative regulatory protein. The prior art also does not teach SE ID NO: 1.  Claims 1-21 meet the criteria set out in PCT Article 33(4), in that they have the industrial applicability in being useful for increasing the methionine content of edible portions of important food crops.  NEW CITATIONS	. statement			
Inventive Step (IS)  Claims  NONE  Claims  Claims  NONE  Claims  Claim 19 lacks novelty under PCT Article 33(2) as being anticipated by Kirihara et al.  Kirihara et al teach the nucleotide sequenc3e of the 10 kDa zein gene from maize. Sequences found within tuntranslated region in the 3' terminus are shown (page 362).  Claim 19 lacks an inventive step under PCT Article 33(3) as being obvious over Kirihara et al.  Kirihara et al is described above. The claims lack an inventive step as the reference describes a product that is encompassed by the claims.  Claims 1-18, 20, and 21 meet the criteria set out in PCT Article 33(2) and (3), because the prior art does not teach or far suggest a DNA construct comprising the coding sequences of a delta-zein operably linked to a 3' UTR that has been modified to no longer contain binding sites for the dzr1 negative regulatory protein. The prior art also does not teach SE ID NO: 1.  Claims 1-21 meet the criteria set out in PCT Article 334), in that they have the industrial applicability in being useful for increasing the methionine content of edible portions of important food crops.	Novelty (N)	Claims	1-18 and 20-21	YE
Industrial Applicability (IA)  Claims  Claims		Claims	19	NC
Industrial Applicability (IA)  Claims  Claims	Inventive Step (IS)	Claims	1-18, 20, and 21	YE
citations and explanations (Rule 70.7)  Claim 19 lacks novelty under PCT Article 33(2) as being anticipated by Kirihara et al.  The claim is broadly drawn towards any isolated nucleic acid comprising the 3' UTR of any 10 kDa zein gene. Kirihara et al teach the nucleotide sequenc3e of the 10 kDa zein gene from maize. Sequences found within t untranslated region in the 3' terminus are shown (page 362).  Claim 19 lacks an inventive step under PCT Article 33(3) as being obvious over Kirihara et al.  Kirihara et al is described above. The claims lack an inventive step as the reference describes a product that is encompassed by the claims.  Claims 1-18, 20, and 21 meet the criteria set out in PCT Article 33(2) and (3), because the prior art does not teach or fa suggest a DNA construct comprising the coding sequences of a delta-zein operably linked to a 3' UTR that has been modified to no longer contain binding sites for the dzr1 negative regulatory protein. The prior art also does not teach SE ID NO: 1.  Claims 1-21 meet the criteria set out in PCT Article 334), in that they have the industrial applicability in being useful for increasing the methionine content of edible portions of important food crops.  NEW CITATIONS		Claims	19	NC
citations and explanations (Rule 70.7)  Claim 19 lacks novelty under PCT Article 33(2) as being anticipated by Kirihara et al.  The claim is broadly drawn towards any isolated nucleic acid comprising the 3' UTR of any 10 kDa zein gene. Kirihara et al teach the nucleotide sequenc3e of the 10 kDa zein gene from maize. Sequences found within tuntranslated region in the 3' terminus are shown (page 362).  Claim 19 lacks an inventive step under PCT Article 33(3) as being obvious over Kirihara et al.  Kirihara et al is described above. The claims lack an inventive step as the reference describes a product that is encompassed by the claims.  Claims 1-18, 20, and 21 meet the criteria set out in PCT Article 33(2) and (3), because the prior art does not teach or fa suggest a DNA construct comprising the coding sequences of a delta-zein operably linked to a 3' UTR that has been modified to no longer contain binding sites for the dzr1 negative regulatory protein. The prior art also does not teach SE ID NO: 1.  Claims 1-21 meet the criteria set out in PCT Article 334), in that they have the industrial applicability in being useful for increasing the methionine content of edible portions of important food crops.	Industrial Applicability (IA)	Claims	1-21	YE
Claim 19 lacks novelty under PCT Article 33(2) as being anticipated by Kirihara et al.  The claim is broadly drawn towards any isolated nucleic acid comprising the 3' UTR of any 10 kDa zein gene. Kirihara et al teach the nucleotide sequenc3e of the 10 kDa zein gene from maize. Sequences found within tuntranslated region in the 3' terminus are shown (page 362).  Claim 19 lacks an inventive step under PCT Article 33(3) as being obvious over Kirihara et al.  Kirihara et al is described above. The claims lack an inventive step as the reference describes a product that is encompassed by the claims.  Claims 1-18, 20, and 21 meet the criteria set out in PCT Article 33(2) and (3), because the prior art does not teach or fail suggest a DNA construct comprising the coding sequences of a delta-zein operably linked to a 3' UTR that has been modified to no longer contain binding sites for the dzr1 negative regulatory protein. The prior art also does not teach SE ID NO: 1.  Claims 1-21 meet the criteria set out in PCT Article 334), in that they have the industrial applicability in being useful for increasing the methionine content of edible portions of important food crops.  NEW CITATIONS	, , ,		MONE	
NONE			s bains abuique quas Visibass at al	

International application No.

PCT/US99/20308

## VIII. Certain observations on the international application

The following observations on the clarity of the claims, description, and drawings or on the question whether the claims are fully supported by the description, are made:

The description is objected to under PCT Rule 66.2(a)(v) as lacking clarity under PCT Article 5 because it fails to contain an adequate written description of all isolated nucleic acids comprising a 3' UTR of all 10 kDa zein genes. The description is inadequate because: the only 3' UTR of a 10 kDa zein gene described by the description is that within the sequence listing. Other such isolated sequences are not described, and therefore not reduced to practice. A description of a nucleic acid required more than just an explanation of a method of how one would obtain it. What is required is a description of the DNA itself.

Claim 19 is objected to as lacking clarity under PCT Rule 66.2(a)(v) because practice of the claimed invention is not adequately described in writing, as required under PCT Rule 5.1(a)(iii), for the reasons set forth in the immediately preceding paragraph.

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Supplemental Box

(To be used when the space in any of the preceding boxes is not sufficient)

Continuation of: Boxes I - VIII

Sheet 10

#### **CLASSIFICATION:**

The International Patent Classification (IPC) and/or the National classification are as listed below: IPC(7): C12N 5/04, 15/29, 15/09, 15/11, 15/63, 15/64, 15/66, 15/67, 15/82: AO1H 5/00, 5/10 and US Cl.: 435/69.I, 320.1, 410, 412, 419, 468; 536/23.6; 800/278, 287, 295, 298, 320.1

### IV. LACK OF UNITY OF INVENTION:

3. This Authority considers that the requirement of unity of invention in accordance with Rules 13.1, 13.2, and 13.3 is not complied with for the following reasons:

As applicant was previously notified this International Preliminary Examining Authority has found plural inventions claimed in the International Application covered by the claims indicated below:

This application contains the following inventions or groups of inventions which are not so linked as to form a single inventive concept under PCT Rule 13.1. In order for all inventions to be examined, the appropriate additional examination fees must be paid.

Group I, claim(s)1-18, drawn to a first product, a DNA construct encoding a delta-zein, linked to a promoter and a 3' untranslated region (UTR) modified to be devoid of dzr1 binding sites, and vectors and plants transformed with said DNA construct; and a first method, comprising making high methionine corn with said DNA construct.

Group II, claim(s) 19-21, drawn to a second product, an isolated nucleic acid comprising a 3' UTR of a 10 kDa zein gene, and a chimeric gene comprising any coding sequence operably linked to said UTR.

and it considers that the International Application does not comply with the requirements of unity of invention (Rules 13.1, 13.2 and 13.3) for the reasons indicated below:

The inventions listed as Groups I and II do not relate to a single inventive concept under PCT Rule 13.1 because, under PCT Rule 13.2, they lack the same or corresponding special technical features for the following reasons: the modified 3' UTR of the first product in Group I is not shared with the non-modified 3' UTR of the second product in Group II. The non-modified UTR of Group II also is no shared or used by the method of Group I. Further, the sequences of both groups may be obtained by alternative means, such as chemical synthesis.